

AMENDMENTS TO THE CLAIMS

This listing of claims will replace all prior versions, and listings, of claims in the application.

1. (Currently amended) A therapeutically effective composition comprising: an isolated and purified antibody (Jianye-2), which is specifically made against the synthetic amino acid sequence, RSATEEEPPNDD (SEO ID NO: 1) and which specifically binds to the amino acid sequence, RSATEEEPPNDD (SEO ID NO: 1) of the α -subunit of the ($\text{Na}^+ + \text{K}^+$)-ATPase enzyme and its isoforms thereof, wherein the binding of the Jianye-2 antibody to the amino acid sequence, RSATEEEPPNDD (SEO ID NO: 1) in a therapeutic amount, increases myocyte intracellular diastolic or systolic calcium, ~~and~~ without inhibiting ($\text{Na}^+ + \text{K}^+$)-ATPase enzyme activity.

2. Cancelled

3. (Currently amended) The therapeutically effective composition of claim 1 wherein binding of the Jianye-2 antibody to the amino acid sequence, RSATEEEPPNDD (SEO ID NO: 1), of the α -subunit of ($\text{Na}^+ + \text{K}^+$)-ATPase in a therapeutic amount, exerts a positive inotropic effect in cardiac myocytes or in a heart for treatment of heart failure and heart muscle contractile disorders.

4. (Currently amended) The therapeutically effective composition of claim 1 wherein the amino acid sequence, RSATEEEPPNDD (SEO ID NO: 1) is an antigen or a component of a vaccine, and wherein the antibody is a polyclonal antibody, a monoclonal antibody, a humanized antibody or a human antibody.

5 – 6 Cancelled

7. (Currently amended) The therapeutically effective composition of claim [[4]] 1 wherein the antibody ~~or antigen or vaccine~~ is administered to a patient in an effective therapeutic amount for treatment of heart failure or heart muscle contractile disorders.

8. (Withdrawn) An antibody which recognizes the amino acid sequence comprising DVEDSYGQQWTYEQR of the .alpha.-subunit of (Na⁺+K⁺)-ATPase enzyme, of which recognizes an isoform of the amino acid sequence.

9. (Withdrawn) The antibody of claim 8, wherein binding of the antibody to the amino acid sequence, DVEDSYGQQWTYEQR, of the .alpha.-subunit of (Na⁺+K⁺)-ATPase increases myocyte intracellular diastolic and systolic calcium.

10. (Withdrawn) The antibody of claim 8, wherein binding of the antibody to the amino acid sequence, DVEDSYGQQWTYEQR, of the .alpha.-subunit of (Na⁺+K⁺)-ATPase exerts a positive inotropic effect in cardiac myocytes.

11. (Withdrawn) The antibody of claim 8, wherein the antibody is a polyclonal antibody.

12. (Withdrawn) The antibody of claim 8, wherein the antibody is a monoclonal antibody.

13. (Withdrawn) The antibody of claim 8, wherein the antibody is a humanized antibody.

14. (Withdrawn) The antibody of claim 8, wherein the antibody is administered to a patient in an effective therapeutic amount to treat the patient suffering from or susceptible to heart disease and/or muscle contractile disorders.

15. (Withdrawn) A purified peptide comprising the amino acid sequence RSATEEEPPNDD or derivatives or isoform thereof.

16. (Withdrawn) The peptide of claim 15, wherein the peptides are administered individually or in combination in a pharmaceutically acceptable carrier to a patient.

17. (Withdrawn) A nucleic acid vector encoding an amino acid sequence comprising RSATEEEPPNDD or isoform thereof.

18. (Withdrawn) The vector of claim 17, wherein the vector comprises tissue specific promoters.

19. (Withdrawn) The vector of claim 17, wherein the tissue specific promoters are cardiac tissue specific.

20. (Withdrawn) The vector of claim 17, wherein the in vivo generated antibodies bind to the amino acid sequence, RSATEEEPPNDD, of the .alpha.-subunit of (Na⁺+K⁺)-ATPase.

21. (Withdrawn) The vector of claim 18, wherein binding of the in vivo generated antibodies to the amino acid sequence, RSATEEEPPNDD, of the .alpha.-subunit of (Na⁺+K⁺)-ATPase increases myocyte intracellular diastolic and systolic calcium.

22. (Withdrawn) The vector of claim 18, wherein binding of the in vivo generated antibodies to the amino acid sequence, RSATEEEPPNDD, of the .alpha.-subunit of (Na⁺+K⁺)-ATPase exerts a positive inotropic effect in cardiac myocytes.

23. (Withdrawn) The vector of claim 18, wherein the vector is administered to a patient in an effective therapeutic amount to treat the patient suffering from or susceptible to heart disease and/or muscle contractile disorders.

24. (Withdrawn) A purified peptide comprising the amino acid sequence DVEDSYGQQWTYEQR or derivative or isoform thereof.

25. (Withdrawn) The peptide of claim 24, wherein the peptides are administered individually or in combination in a pharmaceutically acceptable carrier to a patient.

26. (Withdrawn) A nucleic acid vector encoding an amino acid sequence comprising DVEDSYGQQWTYEQR.

27. (Withdrawn) The vector of claim 26, wherein the vector comprises tissue specific promoters.

28. (Withdrawn) The vector of claim 26, wherein the tissue specific promoters are cardiac tissue specific.

29. (Withdrawn) The vector of claim 26, wherein the in vivo generated antibodies bind to the amino acid sequence, DVEDSYGQQWTYEQR, of the .alpha.-subunit of (Na⁺+K⁺)-ATPase.

30. (Withdrawn) The vector of claim 27, wherein binding of the in vivo generated antibodies to the amino acid sequence, DVEDSYGQQWTYEQR, of the .alpha.-subunit of (Na⁺+K⁺)-ATPase increases myocyte intracellular diastolic and systolic

calcium.

31. (Withdrawn) The vector of claim 27, wherein binding of the in vivo generated antibodies to the amino acid sequence, DVEDSYGQQWTYEQR, of the .alpha.-subunit of (Na⁺+K⁺)-ATPase exerts a positive inotropic effect in cardiac myocytes.

32. (Withdrawn) The vector of claim 27, wherein the vector is administered to a patient in an effective therapeutic amount to treat the patient suffering from or susceptible to heart disease and/or muscle contractile disorders.

33. (Withdrawn) A method of generating antibodies, wherein binding of the antibodies to an epitope of the .alpha.-subunit of (Na⁺+K⁺)-ATPase exerts a positive inotropic effect in cardiac myocytes, comprising: generating amino acid sequences corresponding to overlapping peptide fragments of the .alpha.-subunit of (Na⁺+K⁺)-ATPase and variants thereof; and, obtaining antibodies specific for each peptide fragment by standard methods; and, determining the effects of the antibodies on intracellular diastolic and systolic calcium levels and cell shortenings as compared to controls.

34. (Withdrawn) The method of claim 33, wherein binding of the antibodies to the .alpha.-subunit of (Na⁺+K⁺)-ATPase exerts a positive inotropic effect in cardiac myocytes.

35. (Withdrawn) The method of claim 34, wherein binding of the antibodies to the .alpha.-subunit of (Na⁺+K⁺)-ATPase increases myocyte intracellular diastolic and systolic calcium.

36. (Withdrawn) The method of claim 35, wherein binding of the antibodies to the .alpha.-subunit of (Na⁺+K⁺)-ATPase increases myocytes contractions as compared to controls.

37. (Withdrawn) The method of claim 34, wherein the antibodies generated are polyclonal antibodies.

38. (Withdrawn) The method of claim 34, wherein the antibodies generated are monoclonal antibodies.

39. (Withdrawn) The method of claim 34, wherein the antibody is administered to a patient in an effective therapeutic amount to treat the patient suffering from or susceptible to heart disease and/or muscle contractile disorders.

40. (Withdrawn) The method of claim 34, wherein the antibody is administered to a patient in a therapeutically effective amount to block other molecules from binding to drug-interaction sites of (Na⁺+K⁺)-ATPase, wherein the patient is suffering from or susceptible to arrhythmias, tachyrrhythmias and the like.

41. (Withdrawn) The antibodies of claim 40, wherein the antibodies eliminate negative inotropic agents.

42. (Withdrawn) A method for diagnosing heart failure and/or contractile disorders comprising: isolating heart tissue; and, allowing the binding of inotropic antibodies to epitopes of isolated heart tissue; and, measuring intracellular diastolic and systolic calcium and cell shortenings.

43. (Withdrawn) The method of claim 42, wherein inotropic antibodies binding to epitopes of the .alpha.-subunit of (Na⁺+K⁺)-ATPase from cells of patients suffering from or susceptible to heart failure and/or contractile disorders will have a lower inotropic effect as compared to healthy individuals.

44. (Withdrawn) A method for targeting and blocking the RSATEEEPPNDD site of .alpha.-subunit of the (Na⁺+K⁺)-ATPase, comprising: contacting a myocyte with a desired molecule; and, measuring the intracellular diastolic and systolic Ca²⁺; and, measuring cell shortening and heart function; whereby, identifying molecules useful for therapy of patients suffering from or susceptible to heart disease and other contractile disorders.

45. (Withdrawn) The method of claim 44, wherein the desired molecules are administered to patients suffering from and/or susceptible to heart disease and other contractile disorders, an effective therapeutic amount of desired molecules.

46. (Withdrawn) A method for targeting and blocking the DVEDSYGQQWTYEQR site of .alpha.-subunit of the (Na⁺+K⁺)-ATPase, comprising: contacting a myocyte with a desired molecule; and, measuring the intracellular diastolic and systolic Ca²⁺; and, measuring cell shortening and heart function; whereby, identifying molecules useful for therapy of patients suffering from or susceptible to heart disease and other contractile disorders.

47. (Withdrawn) The method of claim 46, wherein the desired molecules are administered to patients suffering from and/or susceptible to heart disease and other contractile disorders, an effective therapeutic amount of desired molecules.

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REMARKS

Claims 1, 3, 4 and 7 are amended. No new matter is added.

Claims 2, 5 and 6 have been cancelled.

Claims 1, 3, 4 and 7 – 38 are pending.

Claims 1, 3, 4 and 7 are under examination.

1. The withdrawal of the rejection under 35 U.S.C. § 112, 1st paragraph, written description, is noted with appreciation.